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For: Ar

Antisense Modulation of MEKK4 Expression

BOX SEQUENCE Assistant Commissioner for Patents Washington DC 20231

PATENT APPLICATION TRANSMITTAL LETTER

Transmitted herewith for filing, please find the following:

- The specification of the above-referenced patent application;
- An executed Declaration or Oath and Power of Attorney;
- An Assignment of the invention to **Isis Pharmaceuticals Inc**. with recordation cover sheet (PTO Form PTO-1595) and \$40.00 cover fee;
- An executed Verified Statement Claiming Small Entity Status under 37 CFR 1.9 and 1 27:
- Statement to Support Filing and Submission of DNA/Amino Acid Sequences in Accordance with 37 CFR \S \$ 1.821 through 1.825;
- Sequence listing in computer readable form in accordance with 37 C.F.R. § 1.821(e);
- An Information Disclosure Statement with references.

The filing fee has been calculated as shown below:

	Communication Co	250 00 00		
For:	No. Filed	No. Extra	Rate	Fee
BASE FEE			<u> </u>	\$345.00
Total Claims	20 - 20 =	0	X \$9=	\$0
Indep.	2 - 3 =	0	x\$39=	\$0
			TOTAL	\$ 345.00

The Commissioner is hereby authorized to charge the following fees to Deposit Account No. 500252:

- the amount of \$385.00 for the above listed fees;
- payment of the following fees associated with this communication or credit any overpayment;
- any additional filing fees required under 37 CFR 1.16 including fees for presentation of extra claims; and
- ullet any additional patent application processing fees under 37 CFR 1.17 and under 37 CFR 1.20 (d).

Triplicate copies of this transmittal are enclosed.

Date: Sept 29, 2000

Laurel Spear Bernstein Registration No. 37,280 Isis Pharmaceuticals, Inc.

Please address all correspondence to:

Jane Massey Licata Law Offices of Jane Massey Licata 66 East Main Street Marlton NJ 08053 (856) 810-1515 DOCKET NO.:RTS-0169

"Express Mail" Label No.: EL263653993US Date of Deposit: 9-29-00

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Donna T. Ward, William Gaarde, Brett P. Monia and Inventor(s): Jacqueline Wyatt

Serial No.: not yet assigned

herewith Filing Date:

Title: Antisense Modulation of MEKK4 Expression

BOX SEQUENCE Assistant Commissioner for Patents Washington, D.C., 20231

STATEMENT TO SUPPORT FILING AND SUBMISSION IN ACCORDANCE WITH 37 C.F.R. §§ 1.821 THROUGH 1.825

I hereby state, in accordance with the requirements of 37 C.F.R. §1.821(f), that the contents of the paper and computer readable copies of the Sequence Listing, submitted in accordance with 37 C.F.R. §1.821(c) and (e), respectively, are the same.

Date: Sept 29, 2000

Respectfully submitted,

Laurel Spear Bernstein Registration No. 37,280 Isis Pharmaceuticals, Inc.

Please address all correspondence to:

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